TRANSMITTAL LETTER TO THE UNITED STATES 14174-109US1 RIB 016.2PCT DESIGNATED/ELECTED OFFICE (DO/EO/US) U.S. APPLICATION NO. (If know CONCERNING A SUBMISSION UNDER 35 U.S.C. 371 Not Yet Assigned INTERNATIONAL APPLICATION NO. INTERNATIONAL FILING DATE PRIORITY DATE CLAIMED PCT/US2004/001461 21 January 2004 (21.01.2004) 21 January 2003 (21.01.2003) TITLE OF INVENTION Lipophilic Derivatives of Double-Stranded Ribonucleic Acid APPLICANT(S) FOR DO/EO/US Philipp Hadwiger, Matthias John, Christina Lorenz, Hans-Peter Vornlocher and Stefan Limmer Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information: This is a FIRST submission of items concerning a submission under 35 U.S.C. 371. This is a SECOND or SUBSEQUENT submission of items concerning a submission under 35 U.S.C. 371. This is an express request to begin national examination procedures (35 U.S.C. 371(f)). The submission must include items (5), (6), (9) and (21) indicated below. The US has been elected (Article 31). A copy of the International Application as filed (35 U.S.C. 371(c)(2)) is attached hereto (required only if not communicated by the International Bureau). has been communicated by the International Bureau. is not required, as the application was filed in the United States Receiving Office (RO/US). An English language translation of the International Application as filed (35 U.S.C. 371(c)(2)). is attached hereto. has been previously submitted under 35 U.S.C. 154(d)(4). Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3)) are attached hereto (required only if not communicated by the International Bureau). have been communicated by the International Bureau. have not been made; however, the time limit for making such amendments has NOT expired. have not been made and will not be made. An English language translation of amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)). An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)). An English language translation of the annexes to the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371(c)(5)). Items 11 to 20 below concern document(s) or information included: 11. An Information Disclosure Statement under 37 C.F.R. 1.97 and 1.98. 12. An assignment document for recording. A separate cover sheet in compliance with 37 C.F.R. 3.28 and 3.31 is included. 13. A preliminary amendment. 14. An Application Data Sheet under 37 C.F.R. 1.76. A substitute specification. 16. A power of attorney and/or change of address letter. 17. A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 37 C.F.R. 1.821 - 1.825. 18. A second copy of the published international application under 35 U.S.C. 154(d)(4). 19. A second copy of the English language translation of the international application under 35 U.S.C. 154(d)(4). 20. Other items or information: International Search Report; Request for the Recording of a Change under Rule 92bis filed with the International Bureau requesting applicant/inventors Roland Kreutzer and Kathleen Madden Williams be removed. CERTIFICATE OF MAILING BY EXPRESS MAIL EL964754309US Express Mail Label No. July 21, 2005 Date of Deposit:

U.S. APPLICATION NO. (if/known/sec-27 C.F.B. 1.5) Not Cet Assigned 3 4 8 PCT/US2004/001461							ATTORNEY'S DOCKET NUMBER 14174-109US1		
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	CLAIMS		NUMBER FILED NUMBER EXTRA			RATE			
Total Claims		24 - 20 =		4		x \$50	\$200.00		
Independent Claims		4 - 3 =		1		x \$200	\$200.00		
MULTIPLE DEPENDENT CLAIM(S) (if applicable) + \$360							\$0.00		
TOTAL OF ABOVE CALCULATIONS =							\$1,300.00		
Applicant claims small entity status. See 37 C.F.R. 1.27. Fees above are reduced by 1/2.							\$0.00	 	
SUBTOTAL =							\$1,300.00		
Processing fee of \$130 for furnishing the English Translation later than 30 months from the earliest claimed priority date (37 C.F.R. 1.492(i))									
+							\$0.00		
TOTAL NATIONAL FEE =							\$1,300.00		
Fee for recording the enclosed assignment (37 C.F.R. 1.21(h)). The assignment must be accompanied by an appropriate cover sheet (37 C.F.R. 3.28, 3.31). \$40.00 per property +							20.00		
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b. Please	b. Please charge my Deposit Account No. 06-1050 in the amount of \$0.00 to cover the above fees.								
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	The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to Deposit Account No. 06-1050. A duplicate copy of this sheet is enclosed.								
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be included on this form. Provide credit card information and authorization on PTO-2038.									
NOTE: Where an appropriate time limit under 37 C.F.R. 1.495 has not been met, a petition to revive (37 C.F.R. 1.137(a) or (b) must be									
filed and granted to restore the International Application to pending status.									
SEND ALL CORRESPONDENCE TO:							1 / 		
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	PTO	Louis Myers							
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TION RESULT REPORT (JUL. 5. 2005 12)

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E-2) BUSY E-4) NO FACSIMILE CONNECTION

Attorney's Docket No.: 14174-109WO1

OFFICIAL COMMUNICATION

FACSIMULE

FOR THE ATTENTION OF:

THE INTERNATIONAL BUREAU OF WIPO

PCT DIVISION

FAX NO: 011 41 22 338 8740

Applicant Serial No. : Alnylam Europe AG : PCT/US2004/001461

Filing Date

January 21, 2004

Title:

LIPOPHILIC DERIVATIVES OF DOUBLE-STRANDED RIBONUCLEIC

ACIDA

Number of pages including this page

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20 **Switzerland**

Sir:

Attached to this facsimile communication cover sheet is Request for the Recording of a Change under Rule 92bis, faxed this day 5th day of July 2005, to the International Bureau of WIPO.

Respectfully submitted,

Susin C. DiNicola Foreign Patent Paralegal

Fish & Richardson P.C. 225 Franklin Street Boston, MA 02110 Telephone: (617) 542-5070

Fax: (617) 542-8906

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Attorney's Docket No.: 14174-109WO1

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IN THE INTERNATIONAL BUREAU

Applicant: Alnylam Europe AG Serial No.: PCT/US2004/001461 Filed: January 21, 2004

Title : LIF

: LIPOPHILIC DERIVATIVES OF DOUBLE-STRANDED RIBONUCLEIC

ACID

International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20 SWITZERLAND

REQUEST FOR THE RECORDING OF A CHANGE UNDER RULE 92 bis

This is being sent to clarify our earlier Request for the Recording of a Change under Rule 92bis on July 1, 2005, that we wish applicants WILLIAMS, Kathleen M. and KREUTZER, Roland be withdrawn as both applicants and inventors.

Respectfully submitted,

Date: July 5, 2005

Allyson R. Hatton, Ph.D.

Reg. No. 54,154

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Telephone: (617) 542-5070 Facsimile: (617) 542-8906

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CERTIFICATE OF TRANSMISSION BY FACSIMILE

I hereby certify that this correspondence is being transmitted by facsimile to the International Bureau of WIPO on the date indicated below

Date of Transmission

Susan C. DiNicola

Typed or Printed Name of Person Signing Certificate

BEST AVAILABLE COPY